

# Limited Effectiveness of Repeated Delivery of IL18 Adenoviral Vector

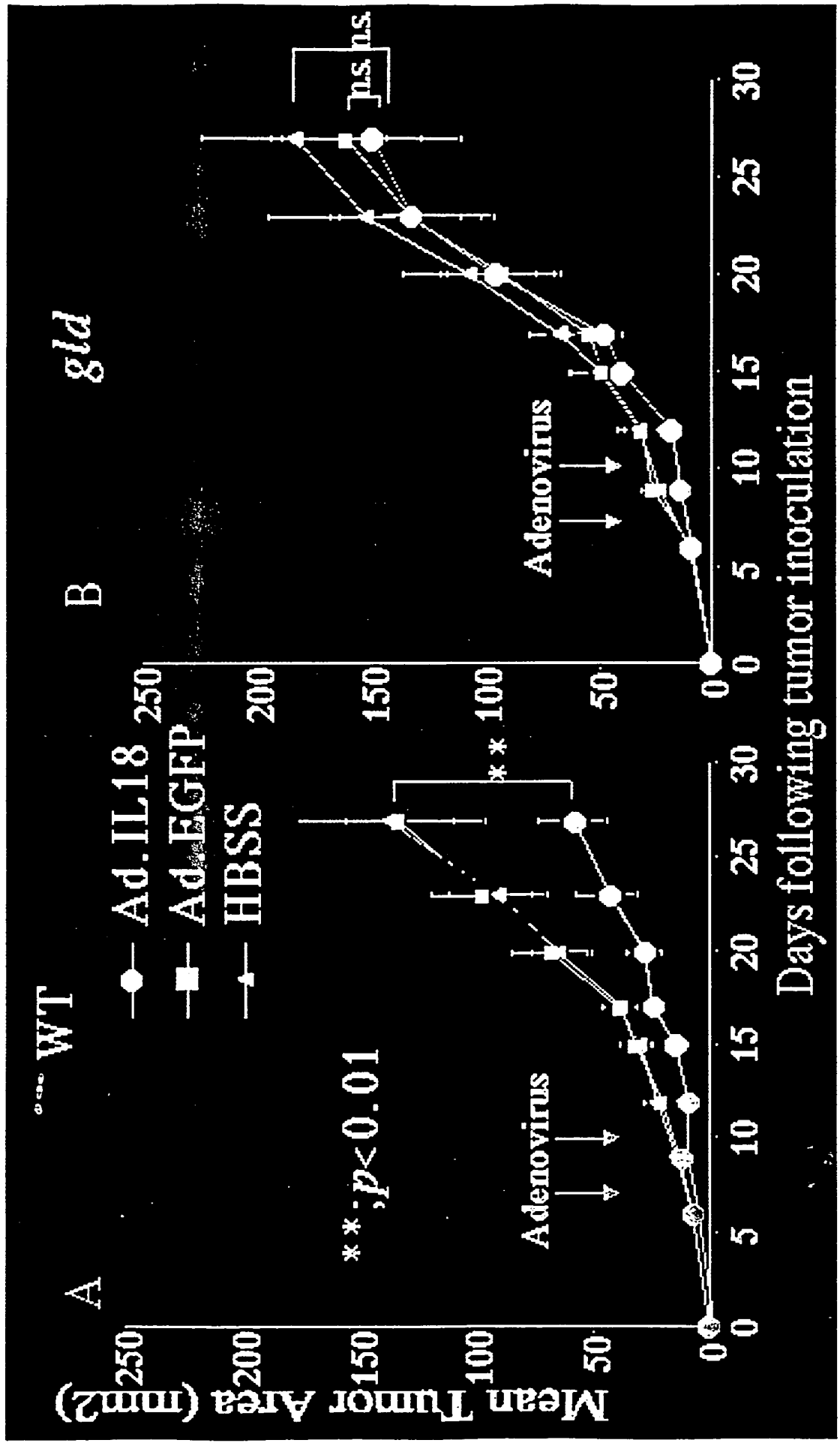


Figure 1A and 1B

# FasL Dependent Induction of Cytotoxicity

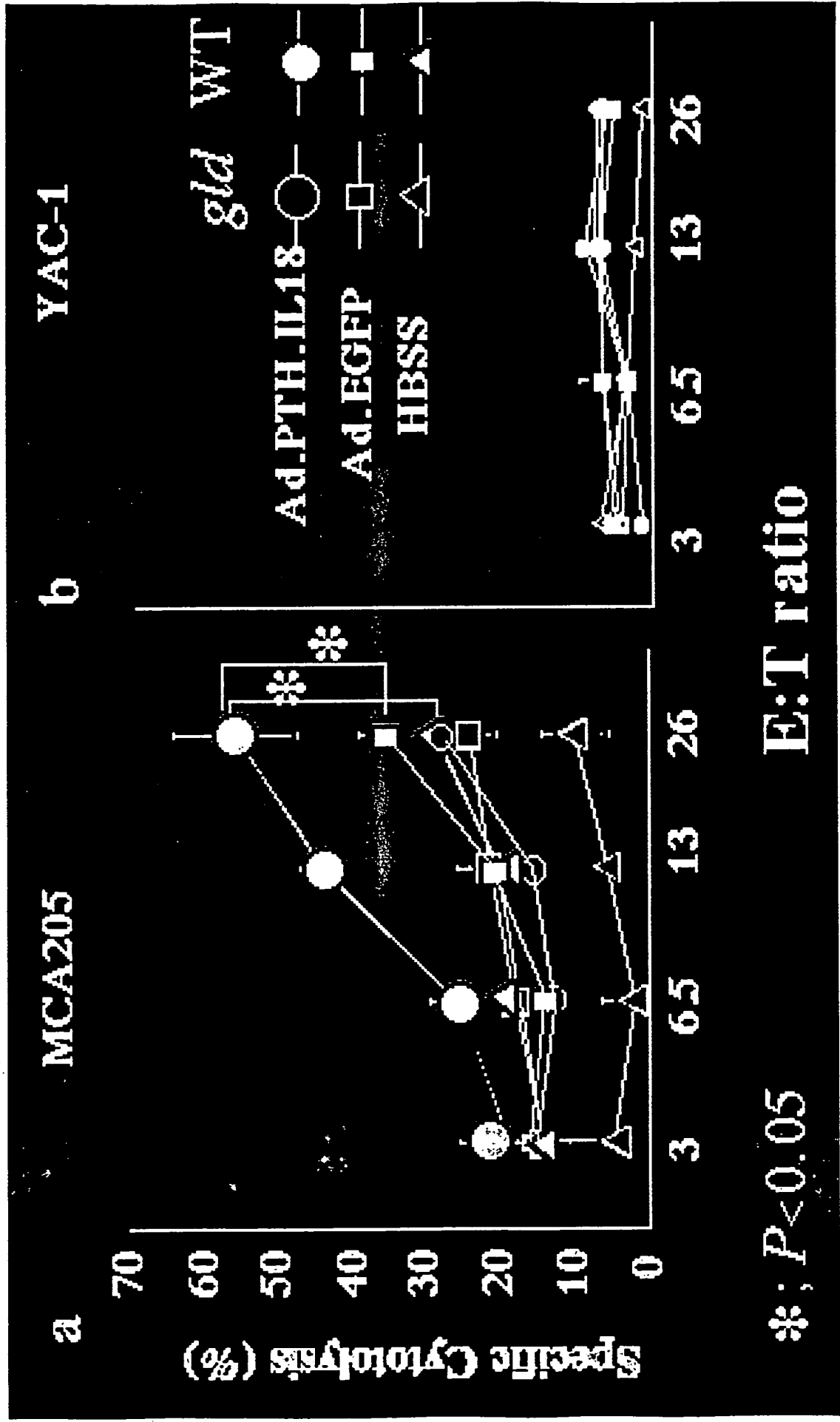


Figure 1c

# DC Combination with Adenoviral Delivery of IL18 Mediates Profound Antitumor Activity in MCA 205

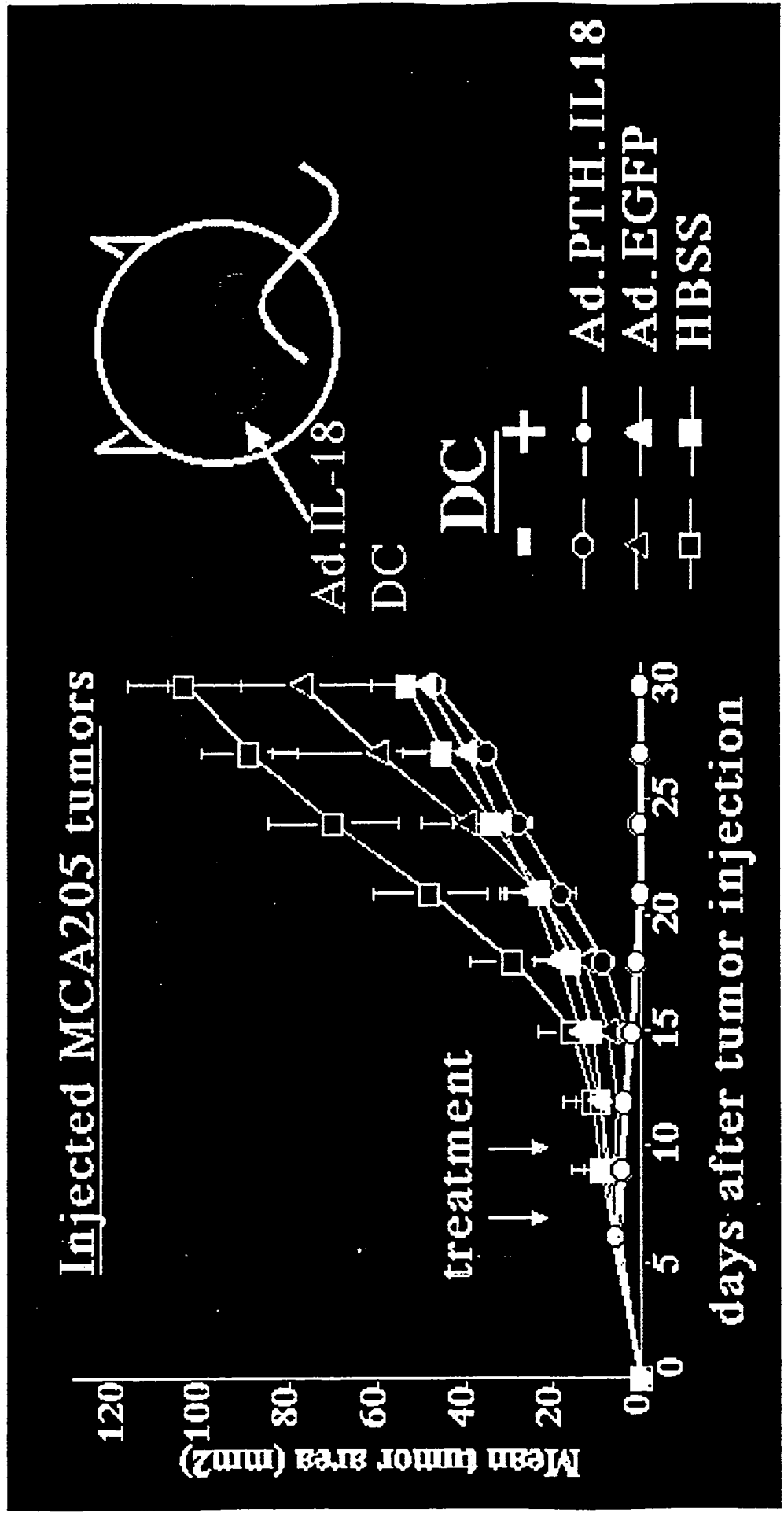


Figure 2A

# Interleukin 18 Mediates Distant Antitumor Effects When Coupled with DC

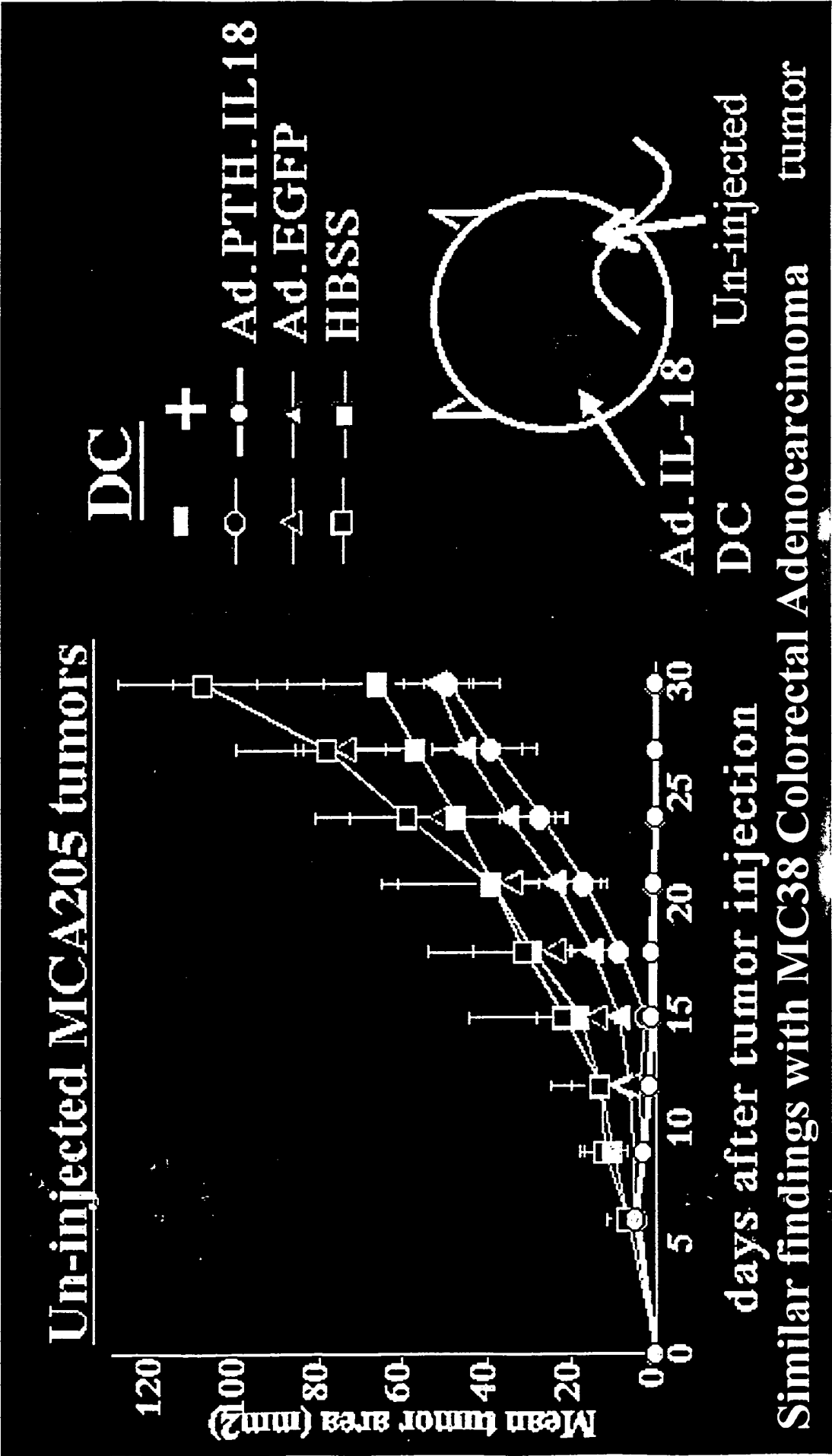


Figure 2B

# Development of Systemic Immunity with IL18 and Dendritic Cell Coadministration

Effector:                      Lymph Node Cells                      Splenocytes

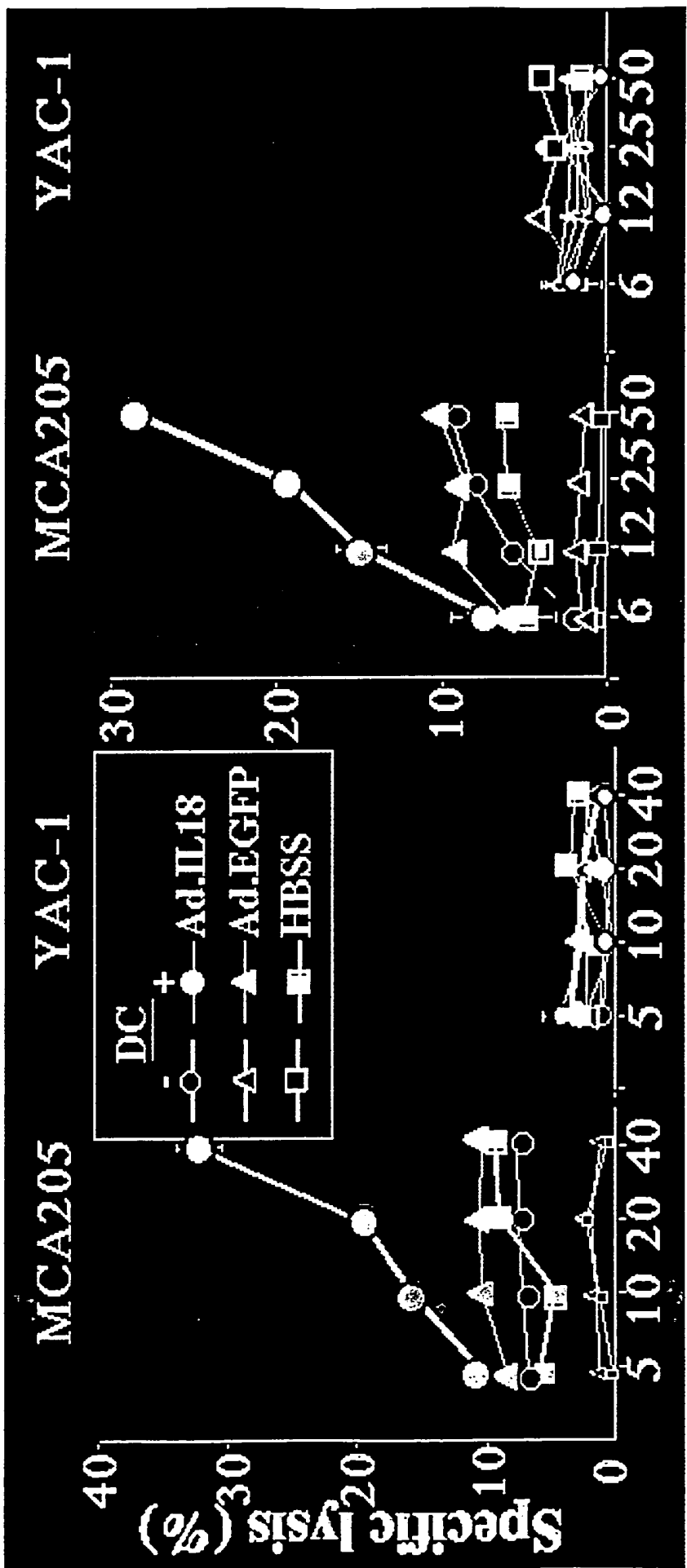


Figure 2C

# Fine Specificity of the Antitumor Cytolytic Response Mediated by IL18 and DCs

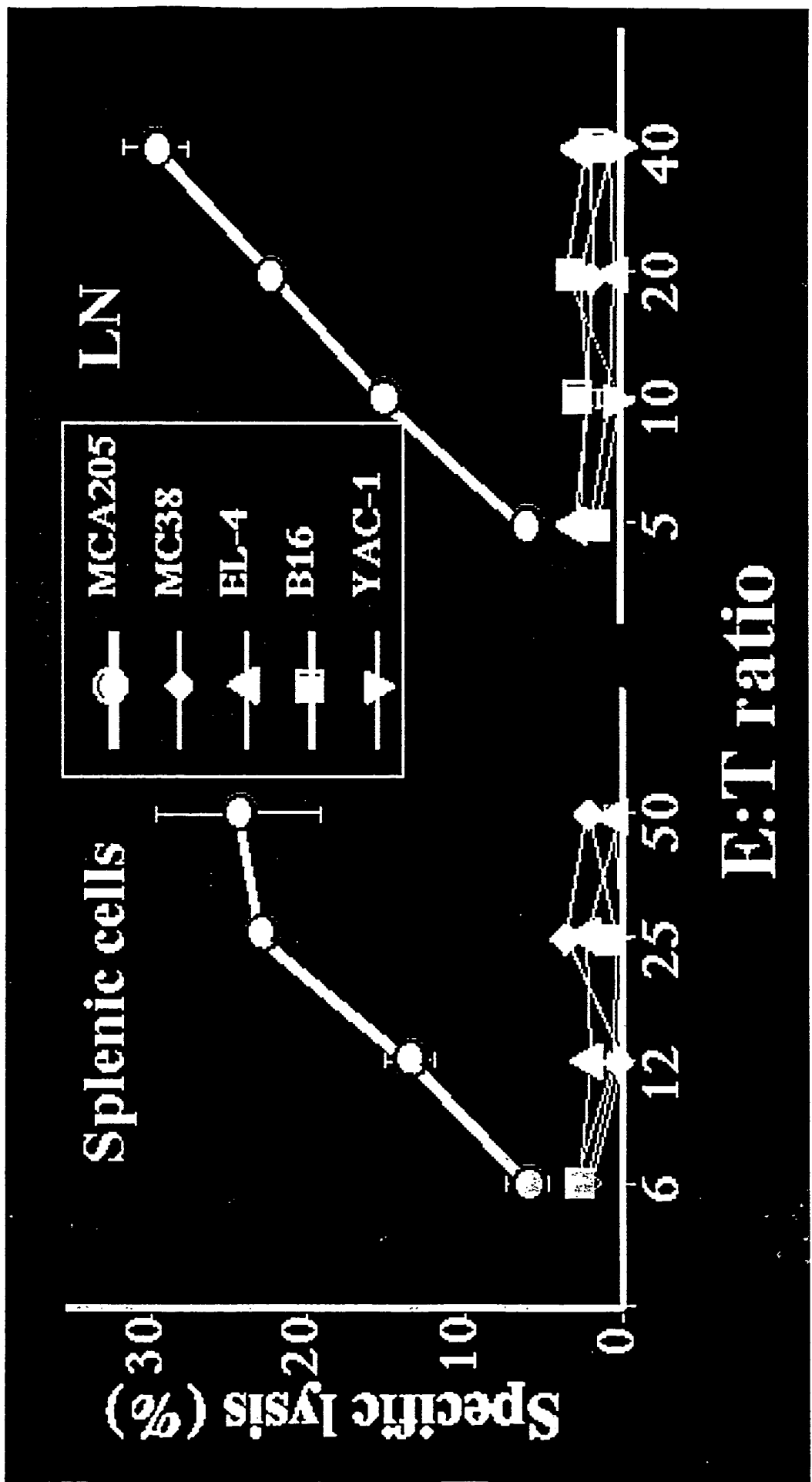


Figure 3A

# Cytolytic Activity Mediated by DC and IL18

## Injection is MHC Class I [K<sup>b</sup>] Restricted

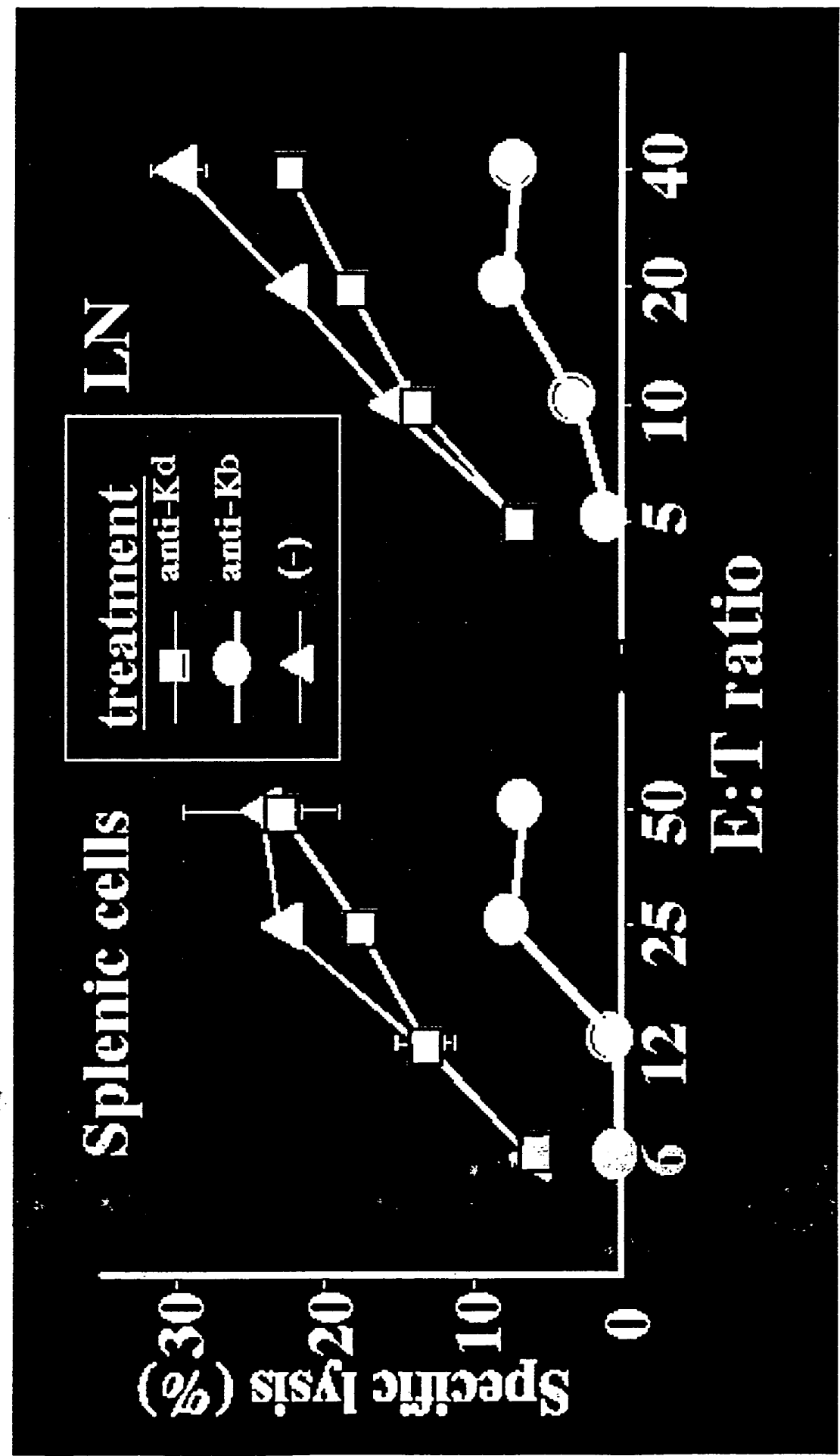


Figure 3B

# DCs Require IL12 Production to Mediate Antitumor Effects with IL18

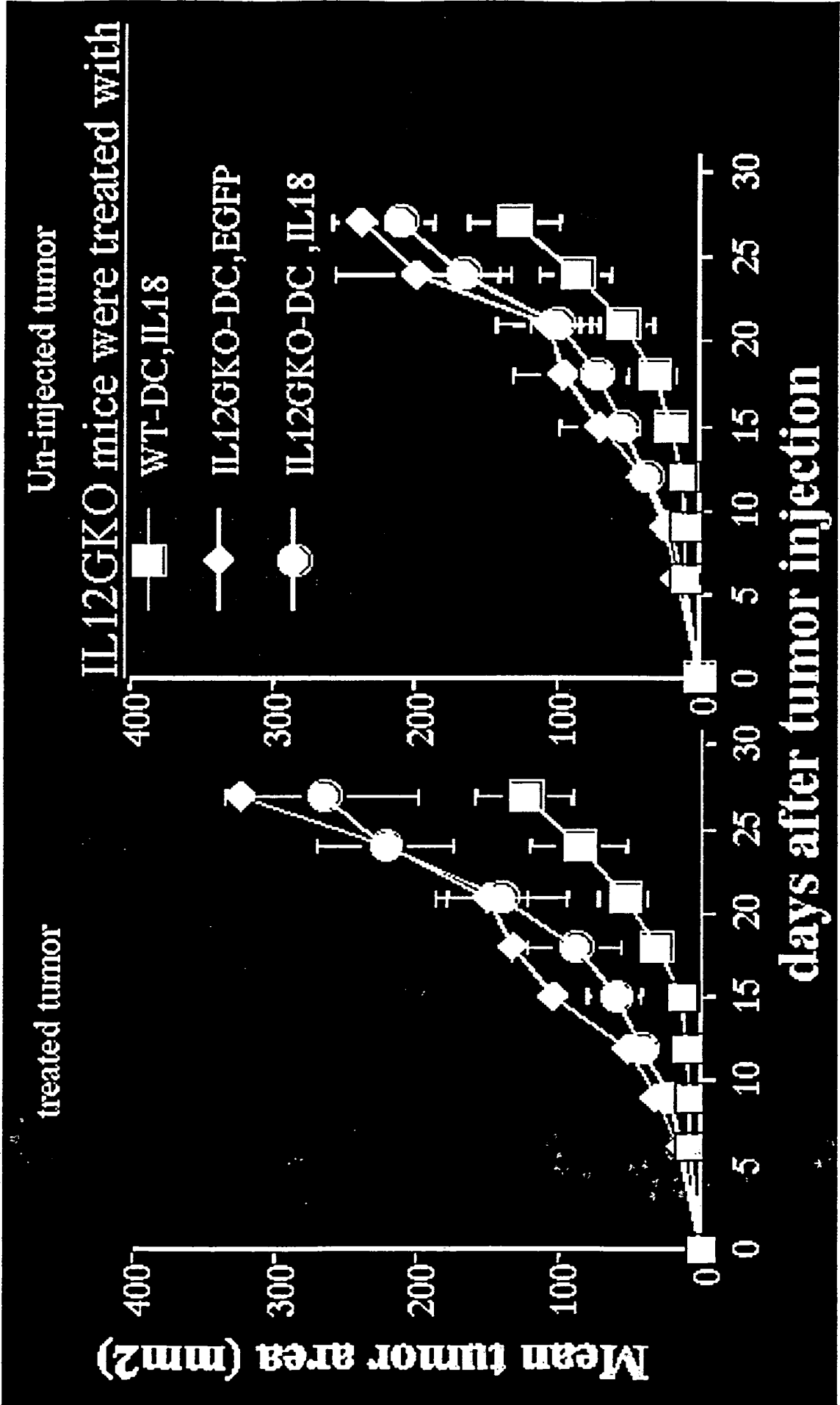


Figure 4



# Nomenclature of novel IL-1 homologues.

Human Gene	Mouse Gene	Protein	Accession Number	old names/Accession (Reference)	Variant/Accession (Reference)
<i>IL1F5<sup>a</sup></i>	<i>Il1f5</i>	IL-1F5	AF186094	IL-1Hy1 / AF186094 (Mulero et al., 1999), FIL1δ / AF201830 (Smith et al., 2000), IL-1H3 / AF200493 (Kumar et al., 2000), IL-1RP3 (Busfield et al., 2000), IL-1L1 AJ242737-38 (Barton et al., 2000), IL-1δ / AF230377 (Debets et al., 2001)	IL-1F5b / AJ242738 (Barton et al., 2000)
<i>IL1F6</i>	<i>Il1f6</i>	IL-1F6	AF201831	FIL1ε / AF201831 (Smith et al., 2000)	
<i>IL1F7</i>	<i>Il1f7<sup>b</sup></i>	IL-1F7	AF201832	FIL1ζ / AF201832 (Smith et al., 2000), IL-1 H4 (Kumar et al., 2000), IL-1RP1 (Busfield et al., 2000), IL-1H AF251118-20 (Pan et al., 2001)	IL-1F7b / AF200496 (Kumar et al., 2000)
					IL-1F7c / AF251120 (Pan et al., 2001)
<i>IL1F8</i>	<i>Il1f8<sup>b</sup></i>	IL-1F8	AF201833	FIL1η / AF201833 (Smith et al., 2000), IL-1H2 / AF200494 (Kumar et al., 2000)	IL-1F8b / AF200494 (Kumar et al., 2000)
<i>IL1F9</i>	<i>Il1f9</i>	IL-1F9	AF200492	IL-1 H1 (Kumar et al., 2000), IL-1RP2 (Busfield et al., 2000), IL-1ε / AF206696 (Debets et al., 2001)	
<i>IL1F10</i>	<i>Il1f10<sup>b</sup></i>	IL-1F10	AF334755	IL-1Hy2 (Lin et al., 2001), FKSG75 (unpublished)	IL-1F10b AY026753 (unpublished)

<sup>a</sup>In this nomenclature IL-1F1, IL-1F2, IL-1F3, and IL-1F4 are reserved for IL-1α, IL-1β, IL-1ra and IL-18, respectively.

<sup>b</sup>A corresponding murine orthologue is yet to be identified.

Figure 5

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IL1a  S-----APFSFLSNVKY-----NFMRIIKYEFILNDALNQSIIRAND--YT 153
IL1b  APV-----RSLNCTLRDSQOKSLVMGSP-----Y-----ELRALPHLOGQD--M-- 152
IL1ra RPSGRKSS-----RMOAERFWQVWORTIFYERNND--VE 39
IL18  -----FGKLESLSVIRNLSDOVLFIDQBNRPFE 67
IL1F5  VLGS-----ALCERMKPSALVLYPHNNQ--IA 28
IL1F6  -----TPQOGSPDISHRNVVLQDOT--IA 32
IL1F7  SFVGENSGVKMGSEDEKDEPQCLEDPAQSPLEPGPSLPTMNFVHTSPKVKNLNPKKSSHDHODHKVVLVDSEN--IA 79
IL1F8  N-----P-----QREA--APSYAHRDSROMHWVLSNS--IA 31
IL1F9  R-----GTPGDADGGGRAV-YQSMC-----KAITGTNDLNGOWMTLOGON--VE 44
IL1F10 C--SLP-----MARYYLKYEADRALYTRDSC--LV 29

IL1a  AA-ALHSLDEAVKFDMCQYK--SKDDAKITVIRISKTQFYVT---AQDESDQVFLKEMPEIPKTTITSETN---LFT 223
IL1b  -----EQOVVFSMSFVOGEESNDKIEVAGLKEKNYPSCVLK--DEKPTLOLESVDPKN-IPKIKMEKF--VE 217
IL1ra GYLQGPNNLESKDVP---IBPH-----ALFEGHCEGKMCISCVY--SEDETRELQEAANNITDISENRKODKE--EAS 107
IL18  DMTDSDCRDNAPRTIFISMKDSQPRGMAVTISVKCEKIST-LSCENK-----IISFKEMPPC---NIKDTSSDIE 137
IL1F5  GGLHSQKVIIGCEHSVVPNRWLDASLS---EVLGGVGGGSOGLSGG---VGOEPTITLSPVNIHELVLAKSSRS--EIE 100
IL1F6  --VPRKDRMSVPLALSSCHVETLEKDRNEVYGLNLNCLMCAH--VGDQPTLOLEKDKIMDLYNQPVPV--SIE 107
IL1F7  --VEDKSYRDEPIFFALSSLSA--SAEKQSP--LGVSKSEFELYCDSDKGQSHESLOLKKEKLMLAAOKESRPPFE 156
IL1F8  --APLSRSKRFVHLHLACRDTFSDKEKNMVYLGIRKDKLFCAE--IQGKPTLOLEKKNIMDLYVERKAOR--PPE 106
IL1F9  --VPRSDSVTVIVAVITCKVPEALBQGRSDFIYLGICNPENCLYCEH--VGEQPTLOLEKKNIMDLYGQPPV--PPE 119
IL1F10 GDPVADNCC-ABKCTLPNGLDRTKV---PFLGIQGGSRCLACVE--TEEGPSLOLEDVNIIEELYKSGGEATE--EIE 101

IL1a  SWETH--STKNYITVEHENLEIAT--KQBY-WPCAGGE-----PSITDFOILEN-----QA 271
IL1b  NKIEI--NNKLEFESAOENNYISTQAENM-PVFLGGTKGGDITDFTMOE-----VSS 269
IL1ra IRSDS--GPTTESAACPQWFLCHAMEADQ-PVSLNNMBDEG--VMVQKPYQED-----E 159
IL18  BQRSVPGHDKMQEESSEYEGYFLACEKEDLFKLKKDELDGDRSIM--FIVONE-----E 193
IL1F5  YSRDM--SLSSSESAAVPCWFLCHVPEADQ-PVRLTQLENGGWNPIITDFYEQC-----E 155
IL1F6  YHSQS--SRNSTESVAFPGWFIIVSSEGGC-PLITTOBLE-----KANTTDGGL-----TMLF 158
IL1F7  YRAQV--GSWNMLESAHPGWFICTSCNCNE-PVGVVDKFE--NRKHIESFGQPVCKAEMSPSEVSD 218
IL1F8  SHNKE--GSTSVROSYSYFGWFIATSTTSGQ-PIFLTKEKE-----ITNNNSYL-----DSVE 157
IL1F9  YRAKT--GRSTLESVAFPGWFIASSRDO--PILLSELS-----KSYNTAFEL-----NIND 169
IL1F10 FOSSS--GSAFRLSAAWPGWFLCGPAPQD-PVQLTKESEPSAR--TKSYTEQS-----W 152

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Figure 6